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DuPont Haskell Laboratory for Health and Environmental Sciences Elkton Road, P.O. Box 50 Newark, DE 19714-0050

Via Federal Express

8EHQ-0305-15944

Document Processing Center (Mail Code 7407M)
Room 6428
Attention: 8(e) Coordinator
Office of Pollution Prevention and Toxics
U.S. Environmental Protection Agency
1201 Constitution Ave., NW
Washington, D.C. 20460

CONTAINS NO US

Dear 8(e) Coordinator:

2,6-Octadienenitrile, 3,7-dimethyl-CAS # 5146-66-7



This letter is to inform you of the preliminary results of a Material Balance and Pharmacokinetics Study in mice that is currently in progress with the above referenced test substance.

As part of the definitive mass balance study, twenty four male and twenty four female mice were dosed with the high dose of the test substance (approximately 600 mg/kg) in corn oil. The dose also contained approximately 900 mg/kg ethanol from the radiolabeled test substance preparation. Following dosing, mice were placed in metabolism cages (4 per cage) for collection of urine and feces. Mice were kept in the metabolism units for 48 hours post dosing then sacrificed by CO<sub>2</sub> asphyxiation. Excreta were collected at 4 time points (6, 12, 24 and 48 hours).

The actual dose rates were  $499.2 \pm 33.1$  mg/kg for males and  $606.3 \pm 18.8$  mg/kg for females at a dose volume of 8 mL/kg. All of the mice appeared lethargic at 2 hours post dosing and normal at 6 hours post dosing. At the 12-hour post dosing collection point, 5 female mice were found dead and 6 more females were sacrificed in extremis. All of the male mice appeared normal. In the mice sacrificed in extremis, the fur was wet, matted and discolored, the animals were cool to the touch, exhibited a greatly reduced ability to move, and showed labored breathing. A brief necropsy of two of the mice sacrificed in extremis and two of the surviving mice found only discoloration in the lungs. No other female mice, or any of the male mice, had clinical signs or mortality at 24 hours. All remaining female mice were sacrificed at 26 hours post dosing.

Under the conditions of this study, the clinical signs observed in the mice are being reported in accordance with the guidance given in the EPA TSCA Section 8(e) Reporting Guide (June 1991).

Sincerely,

A. Michael Kaplan, Ph.D.

AMK/PMH:clp

Director - Regulatory Affairs and Occupational Health

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